

Atty. Dkt. No. SALK3140US-1
(088802-9803)

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Applicants Downes et al.

Title: STRUCTURE OF THE
FARNESOID X RECEPTOR
LIGAND BINDING DOMAIN
AND METHODS OF USE
THEREFOR

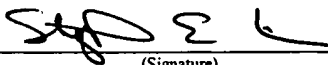
Appl. No.: 10/535,042

Filing Date: 05/13/2005

Examiner: Not yet assigned

Art Unit: 1646

Conf. No. 2218

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| <p align="center">CERTIFICATE OF MAILING</p> <p>I hereby certify that this correspondence is being deposited with the United States Postal Service with sufficient postage as First Class Mail in an envelope addressed to: Commissioner for Patents, P.O. Box 1450, Alexandria, VA 22313-1450, on the date below.</p> <p align="center"><u>Stephen E. Reiter</u> (Printed Name)</p> <p align="center"> (Signature)</p> <p align="center"><u>March 27, 2006</u> (Date of Deposit)</p> |
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INFORMATION DISCLOSURE STATEMENT
UNDER 37 CFR §1.56

Mail Stop Amendment-IDS
Commissioner for Patents
P.O. Box 1450
Alexandria, VA 22313-1450

Sir:

Submitted herewith on Form PTO/SB/08 is a listing of documents known to Applicants in order to comply with Applicants' duty of disclosure pursuant to 37 CFR §1.56.

A copy of each non-U.S. patent document and each non-patent document is being submitted to comply with the provisions of 37 CFR §1.97 and §1.98.

The submission of any document herewith, which is not a statutory bar, is not intended as an admission that such document constitutes prior art against the claims of the present application or that such document is considered material to patentability as defined in 37 CFR

§1.56(b). Applicants do not waive any rights to take any action which would be appropriate to antedate or otherwise remove as a competent reference any document which is determined to be a *prima facie* art reference against the claims of the present application.

TIMING OF THE DISCLOSURE

The listed documents are being submitted in compliance with 37 CFR §1.97(b), before the mailing date of the first Office Action on the merits.

RELEVANCE OF EACH DOCUMENT

All of the documents are in English.

Applicants respectfully request that each listed document be considered by the Examiner and be made of record in the present application and that an initialed copy of Form PTO/SB/08 be returned in accordance with MPEP §609.

The Commissioner is hereby authorized to charge any additional fees which may be required regarding this application under 37 CFR §§ 1.16-1.17, or credit any overpayment, to Deposit Account No. 50-0872. Should no proper payment be enclosed herewith, as by a check being in the wrong amount, unsigned, post-dated, otherwise improper or informal or even entirely missing, the Commissioner is authorized to charge the unpaid amount to Deposit Account No. 50-0872.

Respectfully submitted,

Date 3/27/06

By SEP E.L.

FOLEY & LARDNER LLP
Customer Number: 30542
Telephone: (858) 847-6711
Facsimile: (858) 792-6773

Stephen E. Reiter
Attorney for Applicant
Registration No. 31,192

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| | | | | Application Number | 10/535,042 | |
| | | | | Filing Date | 05/13/2005 | |
| | | | | First Named Inventor | Downes et al. | |
| | | | | Group Art Unit | 1646 | |
| | | | | Examiner Name | Unknown | |
| Sheet | 1 | of | 5 | Attorney Docket Number | | SALK3140US-1 (088802-9803) |

| U.S. PATENT DOCUMENTS | | | | | | |
|-----------------------|-----------------------|----------------------|-----------------------------------|---|---|---|
| Examiner Initials* | Cite No. ¹ | U.S. Patent Document | | Name of Patentee or Applicant of Cited Document | Date of Publication of Cited Document MM-DD-YYYY | Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear |
| | | Number | Kind Code ² (if known) | | | |
| | A1 | 6184353 | | Evans | 02-06-2001 | |

| U.S. PATENT APPLICATION DOCUMENTS | | | | | | |
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| Examiner Initials* | Cite No. ¹ | U.S. Patent Application Document | | Name of Patentee or Applicant of Cited Document | Filing Date of Cited Document MM-DD-YYYY | Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear |
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| FOREIGN PATENT DOCUMENTS | | | | | | | | |
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| | | Office ³ | Number ⁴ | Kind Code ⁵ (if known) | | | | |
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| NON PATENT LITERATURE DOCUMENTS | | | |
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| | A2 | Blumberg and Evans (1998). Orphan nuclear receptors--new ligands and new possibilities. Genes Dev. 12(20), 3149-55. | |
| | A3 | Blumberg et al. (1998). SXR, a novel steroid and xenobiotic-sensing nuclear receptor. Genes Dev. 12(20), 3195-3205. | |
| | A3 | Chiang (2002) Bile Acid regulation of gene expression: roles of nuclear hormone receptors. Endocr Rev. 23(4), 443-463. | |

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*EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant.

¹ Unique citation designation number. ² See attached Kinds of U.S. Patent Documents. ³ Enter Office that issued the document, by the two-letter code (WIPO Standard ST.3). ⁴ For Japanese patent documents, the indication of the year of the reign of the Emperor must precede the serial number of the patent document. ⁵ Kind of document by the appropriate symbols as indicated on the document under WIPO Standard ST. 16 if possible. ⁶ Applicant is to place a check mark here if English language Translation is attached.

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| | A4 | Egea et al. (2000). Crystal structure of the human RXRa ligand-binding domain to its natural ligand: 9-cis retinoic acid EMBO J. 19, 2592-2601. | | |
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| | A11 | Laffitte et al. (2000). Identification of the DNA binding specificity and potential target genes for the farnesoid X-activated receptor. J Biol Chem. 275(14), 10638-47 | | |

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| | A25 | Wang et al. (1999) Endogenous bile acids are ligands for the nuclear receptor FXR/BAR. Mol Cell. 3(5), 543-53 | |

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| | A26 | Watkins et al. (2001). The Human Nuclear Xenobiotic Receptor PXR: Structural Determinants of Directed Promiscuity, Science, 292, 2329-2333 | |
| | A27 | Xu et al. (2001). Structural determinants of ligand binding selectivity between the peroxisome proliferator-activated receptors. Proc Natl Acad Sci U S A. 98(24), 13919-24 | |

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